№ expression or activity [of a daf-18 gene], comprising:

- (a) providing a [cell] <u>nematode</u>, <u>isolated nematode cell</u>, <u>or isolated mammalian cell</u> expressing a <u>nematode</u> <u>daf-18</u> gene; and
- (b) contacting said [cell] <u>nematode</u>, isolated <u>nematode</u> cell, or isolated <u>mammalian</u> cell with a candidate compound, an alteration in *daf-18* expression or activity following contact of said nematode, isolated <u>nematode</u> cell, or isolated <u>mammalian</u> cell with said candidate compound identifying a modulatory compound.

(Amended) A method for identifying a compound that modulates [the] <u>PTEN</u> expression or activity [of a *daf-18* gene], comprising:

- (a) providing a [cell] <u>nematode or isolated nematode cell</u> comprising a mutation in [a] <u>its endogenous</u> *daf-18* gene;
- (b) expressing in said [cell] <u>nematode or isolated nematode cell</u> a mammalian [DAF-18 homologue] <u>PTEN gene</u>; and
- (c) contacting said [cell] <u>nematode or isolated nematode cell</u> with a candidate compound, an alteration in [said mammalian DAF-18] <u>PTEN</u> expression or activity following contact with said candidate compound identifying a modulatory compound.
- 3. (Amended) The method of claim 1 or 2, wherein said compound increases *daf-*18 or PTEN expression or activity and is therefore [capable of] a candidate compound for

increasing longevity of a cell or organism.

Cont

4. (Amended) The method of claim 1 or 2, wherein said compound decreases daf18 or PTEN expression or activity and is [capable of] therefore a candidate compound for treating an impaired glucose tolerance condition or obesity.

ca

8. (Twice Amended) The method of claim [3] 2, wherein said [DAF-18 homologue] PTEN is [a] human [homologue] PTEN.

13

- 10. (Amended) A method for identifying a compound that is [capable of] a candidate compound for ameliorating or delaying an impaired glucose tolerance condition or obesity, comprising contacting a biological sample with a candidate compound and assaying said sample for DAF-18-mediated lipid phosphatase activity, a decrease in said activity indicating a candidate compound [capable of] for ameliorating or delaying an impaired glucose tolerance condition or obesity.
- 11. (Amended) A method for identifying a compound that is [capable of] a candidate compound increasing longevity of a cell or organism, comprising contacting a biological sample with a candidate compound and assaying said sample for DAF-18-mediated lipid phosphatase activity, an increase in said activity indicating a candidate

compound [capable of] for increasing longevity of a cell or organism.

- 12. (Amended) A method for identifying a compound that is [capable of] a candidate compound for ameliorating or delaying an impaired glucose tolerance condition or obesity, comprising contacting a biological sample with a candidate compound and assaying said sample for PTEN-mediated lipid phosphatase activity, a decrease in said activity indicating a candidate compound [capable of] for ameliorating or delaying an impaired glucose tolerance condition or obesity.
- 13. (Amended) A method for identifying a compound that is [capable of] a candidate compound for increasing longevity of a cell or organism, comprising contacting a biological sample with a candidate compound and assaying said sample for PTEN-mediated lipid phosphatase activity, an increase in said activity indicating a candidate compound [capable of] for increasing longevity of a cell or organism.
- 14. (Twice Amended) The method of claim 10 or 12, wherein said method further comprises assaying said compound in a [cell] nematode or isolated nematode cell which comprises a mutation in [a] its endogenous daf-18 gene and which expresses a mammalian [DAF-18 homologue] PTEN gene, a decrease in [DAF-18] PTEN activity indicating a candidate compound [capable of] for treating an impaired glucose tolerance

condition or obesity.

15. (Twice Amended) The method of claim 11 or 13, wherein said method further comprises assaying said compound in a [cell] nematode or isolated nematode cell which comprises a mutation in [a] its endogenous daf-18 gene and which expresses a mammalian [DAF-18 homologue] PTEN gene, an increase in [DAF-18] PTEN activity indicating a candidate compound [capable of] for increasing longevity of a cell or organism.

MNY.

- 16. (Twice Amended) The method of claim 14, wherein said mammalian [DAF-18 homologue] <u>PTEN</u> is human PTEN.
- 17. (Twice Amended) The method of claim 15, wherein said mammalian [DAF-18 homologue] <u>PTEN</u> is human PTEN.

C4

25. (Twice Amended) The transgenic nematode of claim 23, wherein said nematode carries a mutation in [a] <u>its endogenous</u> *daf-18* gene.

Add the following new claims 27 and 28.

(5

--27. The transgenic nematode of claim 23, wherein said mammalian PTEN is